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**INFORMATION DISCLOSURE
STATEMENT BY APPLICANT**

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| Application Number | 10/701,013 |
| Filing Date | 11/04/2003 |
| First Named Inventor | Rory A.J. Curtis |
| Art Unit | 1647 |
| Examiner Name | Allen, Marianne P |
| Attorney Docket Number | 004974.01183 |

Sheet 1 of 15

OTHER PRIOR ART -- NON PATENT LITERATURE DOCUMENTS

| Examiner Initials* | Cite No. ¹ | Include the name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume/issue number(s), publisher, city and/or country where published. | 7 ² |
|--------------------|-----------------------|---|----------------|
| /MA/ ↓ | C1 | Database EMBL Accession No. AA553891, "Homo Sapiens cDNA Clone IMAGE: 1015480 3', mRNA Sequence." Strausberg, Robert (1997-09-11). | |
| | C2 | Database EMBL Accession No. A1740592, "Homo Sapiens cDNA Clone IMAGE: 2365922 3', mRNA Sequence." Strausberg, Robert (1998-06-28). | |
| | C3 | Database EMBL Accession No. AQ310967, "Homo Sapiens Genomic Clone 2384B13, Genomic Survey Sequence." Adams, M. D. et al. (1998-12-23). | |
| | C4 | Janeway, C. A. Jr., and Travers, P., Immunobiology: The Immune System in Health and Disease, Current Biology Ltd/Garland Publishing Inc., London, UK and New York, NY, USA, 1994, p. G-1. | |
| | C5 | Arrighi, I. et al., "Structure, Chromosome Localization, and Tissue Distribution of the Mouse TWIK K ⁺ Channel Gene," FEBS Lett. 425(2):310-316 (1998) | |
| | C6 | Bonaldo, M.F., et al., "Normalization and Subtraction: Two Approaches to Facilitate Gene Discovery," Genome Res. 6(9):791-806 (1996) | |
| | C7 | Cross, S. H. et al., "Purification of CpG Islands Using a Methylated DNA Binding Column," Nat. Genet. 6(3):236-244 (1994) | |
| | C8 | DiPolo, A. et al., "Isolation and Initial Characterization of the 5' Flanking Region of the Human and Murine Cyclic Guanosine Monophosphate-Phosphodiesterase β -Subunit Genes," Invest. Ophthalmol. Vis. Sci. 37(4):551-560 (1996) | |
| | C9 | Doyle, D. A., et al., "The Structure of the Potassium Channel: Molecular Basis of K ⁺ Conduction and Selectivity," Science 280:69-77 (1998) | |

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| /MA/ ↓ | C11 | Fink, M. et al., "Cloning, Functional Expression and Brain Localization of a Novel Unconventional Outward Rectifier K ⁺ Channel," EMBO J. 15(24):6854-6862 (1996) | |
| | C12 | Fink, M. et al., "A Neuronal Two P Domain K ⁺ Channel Stimulated by Arachidonic Acid and Polyunsaturated Fatty Acids," EMBO J. 17(12):3287-3308 (1998) | |
| | C13 | Goldstein, S. A. et al., "ORK1, a Potassium-Selective Leak Channel with Two Pore Domains Cloned from <i>Drosophila Melanogaster</i> by Expression in <i>Saccharomyces Cerevisiae</i> ," PHAS USA 93(23):13256-13261 (1996) | |
| | C14 | Goldstein, S. A. et al., "Sequence and Function of the Two P Domain Potassium Channels: Implications of an Emerging Superfamily," J. Mol. Med. 76:13-20 (1998) | |
| | C15 | Ketchum, K. A. et al., "A New Family of Outwardly Rectifying Potassium Channel Proteins with Two Pore Domains in Tandem," Nature 376(6542):690-695 (1995) | |
| | C16 | Leonoudakis, D. et al., "An Open Rectifier Potassium Channel with Two Pore Domains in Tandem Cloned from Rat Cerebellum," J. Neuroscience 18(3):868-877 (1998) | |
| | C17 | Lesage, F. et al., "Dimerization of TWIK-1 K ⁺ Channel Subunits Via a Disulfide Bridge," EMBO J. 15(23):6400-6407 (1996) | |
| | C18 | Lesage, F. et al., "TWIK-1, a Ubiquitous Human Weakly Inward Rectifying K ⁺ Channel with a Novel Structure," EMBO J. 15(5):1004-1011 (1996) | |
| | C19 | Lesage, F. et al., "The Structure, Function and Distribution of the Mouse TWIK-1 K ⁺ Channel," FEBS Lett. 402(1):28-32 (1997) | |

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| /MA/ | C20 | Martin Gallardo, A. et al., "Automated DNA Sequencing and Analysis of 106 Kilobases from Human Chromosome 19q13.3," Nature Genet. 1:34-30 (1992) | |
| | C21 | Miosga, T. et al., "Sequence and Function analysis of a 9.46 kb Fragment of Saccharomyces Cerevisiae Chromosome X," Yeast 10(7):965-973 (1994) | |
| | C22 | Pinaud, E. et al., "Identification of a Homolog of the C Alpha 3'/hs3 enhancer and of an Allelic Variant of the 3'IGH/hs1,2 Enhancer Downstream of the Human Immunoglobulin Alpha 1 Gene," Eur. J. Immunol. 27(11):2981-2985 (1997) | |
| | C23 | Reid, J. D. et al., "The S. Cerevisiae Outwardly-Rectifying Potassium Channel (DUK1) Identifies a New Family of Channels with Duplicated Pore Domains," Recept. Channels 4(1):51-62 (1996) | |
| | C24 | Royes, R. et al., "Cloning and Expression of a Novel pH-Sensitive Two Pore Domain K ⁺ Channel from Human Kidney," J. Biol. Chem. 273(47):30863-30869 (1998) | |
| | C25 | Wilson, R. et al., "2.2 Mb of Contiguous Nucleotide Sequence from Chromosome III of C. Elegans," Nature 368(6466):32-38 (1994) | |
| | C26 | Yamaguchi, M. et al., "The Primary Structure of the Rat Guanylyl Atrial Natriuretic Peptide Receptor Gene," J. Biol. Chem. 265(33):20414-20420 (1990) | |
| | C27 | GenBank Accession No. AA925138 for UI-R-A1-en-b-02-0-UI.s1 UI-R-A1 Rattus Norvegicus cDNA Clone UI-R-A1-en-b-02-0-UI 3' similar to gi: 2809390; gb: AF031384 Rattus Norvegicus TWIK-related acid Sensitive K ⁺ channel (TASK) mRNA, complete cds, mRNA sequence, February 7, 1999 | |
| | C28 | GenBank Accession No. AAA81455 for Similar to Potassium Channel Proteins and to C. Elegans Protein F2287.7; November 14, 1995 | |

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|--------------------|-----------------------|---|----------------|
| /MA/ ↓ | C29 | GenBank Accession No. AAA83199 for Weak Similarity to a Short Region of Putative Potassium Channel Proteins; December 12, 1985 | |
| | C30 | GenBank Accession No. AAA86127 for Coded for by C. Elegans cDNA yk9g1.3; coded for by C. Elegans cDNA yk8g1.5; coded for by C. Elegans cDNA CEESU55F; April 18, 1996 | |
| | C31 | GenBank Accession No. AAB01688 for TWIK-1, June 7, 1996 | |
| | C32 | GenBank Accession No. AAB04567 for Weak Similarity to a Potassium Channel Protein, July 22, 1996 | |
| | C33 | GenBank Accession No. AAB52940 for Weak Similarity to S. Cerevisiae Outward-Rectifier Potassium Channel TOK1, April 21, 1997 | |
| | C34 | GenBank Accession No. AAB54033 unidentified protein, May 12, 1997 | |
| | C35 | GenBank Accession No. AAB61602 for rakCNK1, June 18, 1997 | |
| | C36 | GenBank Accession No. AAC16973 for TWIK-1 K ⁺ Channel, September 29, 1998 | |
| | C37 | GenBank Accession No. AAC32854 for Putative Potassium Channel Subunit n2P17m1-2, December 21, 1998 | |

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| /MA/ | C38 | GenBank Accession No. AAC32855 for Putative Potassium Channel Subunit n2P17m1-1, December 21, 1998 | |
| | C39 | GenBank Accession No. AAC32856 for Putative Potassium Channel Subunit n2P16, December 21, 1998 | |
| | C40 | GenBank Accession No. AAC32857 for Putative Potassium Channel Subunit n2P20, December 21, 1998 | |
| | C41 | GenBank Accession No. AAC32858 for Putative Potassium Channel Subunit n2P17m2-2, December 21, 1998 | |
| | C42 | GenBank Accession No. AAC32859 for Putative Potassium Channel Subunit n2P17m2-3 | |
| | C43 | GenBank Accession No. AAC32860 for Putative Potassium Channel Subunit n2P17m3, December 21, 1998 | |
| | C44 | GenBank Accession No. AAC32861 for Putative Potassium Channel Subunit n2P18, December 21, 1998 | |
| | C45 | GenBank Accession No. AAC32862 for Putative Potassium Channel Subunit n2P17m2-1, December 21, 1998 | |
| ↓ | C46 | GenBank Accession No. AAC32863 for Putative Potassium Channel Subunit n2P38, December 21, 1998 | |

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No. 7

GenBank Accession No. AAC32865 for Putative Potassium Channel Subunit $\alpha 2P24$; December 21, 1998

/MA/

C48

GenBank Accession No. AAC39952 for TWIK-related acid-sensitive K⁺ channel, February 2, 1999

C48

GenBank Accession No. AAC40181 for TRAAK K+ channel subunit, February 2, 1999

C50

GenBank Accession No. AAC47976 for Contains Similarity to K+ channel proteins, February 26, 1998

CS1

GenBank Accession No. AAC48050 for Similar *S. cerevisiae* Outward-Rectified Potassium channel TOK1, June 2, 1999

CS2

GenBank Accession No. AAC51777 for TWIK-related acid-sensitive K⁺ channel, February 4, 1999

C53

GenBank Accession No. AAC53005 for TREK-1 K+ channel Subunit, February 4, 1990

C54

GenBank Accession No. AAC53367 for TWIK-related acid-sensitive K⁺ channel, February 4, 1999

C55

GenBank Accession No. AAC69250 for Two P domain Potassium channel ORK1, October 29, 1998

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| IMA/  | C56 | GenBank Accession No. AAC71141 for Proline-rich, November 5, 1998 | |
| | C57 | GenBank Accession No. AAC71151 for Contains similarity to outward-rectified potassium channels, November 5, 1998 | |
| | C58 | GenBank Accession No. AAC79458 for Two-pore domain K ⁺ channel TASK-2, November 17, 1998 | |
| | C59 | GenBank Accession No. AAD01203 for Two-pore domain channel TPKC1, July 2, 1997 | |
| | C60 | GenBank Accession No. AAD09336 for Putative potassium channel TWIK, January 26, 1999 | |
| | C61 | GenBank Accession No. AAD09337 for Putative potassium channel DP3, January 26, 1999 | |
| | C62 | GenBank Accession No. AAD09338 for Putative potassium channel DP4, January 26, 1999 | |
| | C63 | GenBank Accession No. AB008537 for Mus Musculus mRNA for cTBAK, complete cds, February 13, 1999 | |
| | C64 | GenBank Accession No. AB013345 for Mus Musculus mRNA for cTBAK, complete cds, May 21, 1998 | |

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| /MA/ ↓ | C65 | GenBank Accession No. AC005035 for Homo Sapiens BAC clone NH0353P23 from 2, complete sequence, December 3, 1998. | |
| | C66 | GenBank Accession No. AC005183 for Homo Sapiens Chromosome 12p13.3, August 21, 1998 | |
| | C67 | GenBank Accession No. AC005290 Unidentified protein, December 10, 1998 | |
| | C68 | GenBank Accession No. AC005848 for Homo Sapiens Chromosome 11 clone CIT987SK-1012F4, October 22, 1998 | |
| | C69 | GenBank Accession No. AC006126 for Homo Sapiens Chromosome 19, cosmid F18718, December 17, 1998 | |
| | C70 | GenBank Accession No. AC006167 for Homo Sapiens clone UWGC.y67c092 from 6p21, December 8, 1998 | |
| | C71 | GenBank Accession No. AF004695 for Oryctolagus cuniculus double-pore potassium channel rabKCNK1 mRNA, June 25, 1997 | |
| | C72 | GenBank Accession No. AF006824 for Mus Musculus TWIK-related acid-sensitive K ⁺ channel (TASK) mRNA, October 7, 1997 | |
| | C73 | GenBank Accession No. AF022821 for Mus Musculus putative potassium channel DP4 mRNA, January 26, 1999 | |

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Date

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**INFORMATION DISCLOSURE
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Sheet 9 of 15

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| | |
|------------------------|-------------------|
| Application Number | 10/701,013 |
| Filing Date | 11/04/2003 |
| First Named Inventor | Rory A.J. Curtis |
| Art Unit | 1647 |
| Examiner Name | Allen, Marianne P |
| Attorney Docket Number | 004974.01183 |

OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS

| Examiner Initials* | Cite No. ¹ | Include the name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published. | 1 ² |
|--------------------|-----------------------|---|----------------|
| /MA/ | C74 | GenBank Accession No. AF033017 for Mus Musculus TWIK-1 K ⁺ channel mRNA, May 23, 1998 | |
| | C75 | GenBank Accession No. AF058492 for Mus Musculus TRAAK K ⁺ channel subunit mRNA, July 21, 1998 | |
| | C76 | GenBank Accession No. AF084830 for Homo Sapiens two pore domain K ⁺ channel (TASK2) mRNA, November 25, 1998 | |
| | C77 | GenBank Accession No. AJ001356 for Orchis morio (Orchidaceae) microsatellite, September 16, 1997. | |
| | C78 | GenBank Accession No. AQ240175 for CIT-HSP-2385B13.TR.1 CIT-HSP Homo Sapiens genomic clone 2385B13, September 29, 1998 | |
| | C79 | GenBank Accession No. AQ319945 for RPC111-111L8.TJ RPC1-11 Homo Sapiens genomic clone. RPC1-11-111L8, May 5, 1999 | |
| | C80 | GenBank Accession No. B82804 for RPC111-17N16.TP RPC1-11 Homo Sapiens genomic clone RPC1-11-17N16, April 8, 1999 | |
| | C81 | GenBank Accession No. BAA25436 for cTBAK, February 13, 1999 | |
| | C82 | GenBank Accession No. CAA19494 for Similar to potassium channel protein, December 18, 1998 | |

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| Examiner Signature | /Marianne Allen/ | Date Considered | 06/26/2007 |
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Sheet 10 of 15

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| Application Number | 10/701,013 |
| Filing Date | 11/04/2003 |
| First Named Inventor | Rory A.J. Curtis |
| Art Unit | 1647 |
| Examiner Name | Allen, Marianne P |
| Attorney Docket Number | 004974.01183 |

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|--------------------|-----------------------|--|----------------|
| /MA/ | C83 | GenBank Accession No. CAA21041 for cDNA EST yk449a10.5 comes from this gene; cDNA EST EMBL: D35319 comes from this gene; cDNA EST EMBL: c12322 comes from this gene; cDNA EST EMBL: D32633 comes from this gene; November 23, 1998 | |
| | C84 | GenBank Accession No. CAA21749 for Predicted using Genefinder; cDNA EST EMBL: Z14528 comes from this gene; November 23, 1998 | |
| | C85 | GenBank Accession No. CAA64176 for Outward-rectifier potassium channel; February 23, 1996 | |
| | C86 | GenBank Accession No. CAA84797 for Weak similarity with Drosophila Melanogaster potassium channel proteins; November 21, 1998 | |
| | C87 | GenBank Accession No. CAA90083 for Weak similarity to a protein domain (segment H5) thought to line the channel pore of potassium channels; cDNA EST EMBL: T00465 comes from this gene; cDNA EST EMBL: D36509 comes from this gene; November 23, 1998 | |
| | C88 | GenBank Accession No. CAA90259 for M110.2; November 23, 1998 | |
| | C89 | GenBank Accession No. CAA91376 for Similarity to potassium channel protein; cDNA EST EMBL: D70075 comes from this gene; cDNA EST EMBL: D66354 comes from this gene; November 23, 1998 | |
| | C90 | GenBank Accession No. CAA92504 for Similarity to Drosophila potassium channel protein (PIR Acc. No. JH0697); cDNA EST EMBL: M89234 comes from this gene; December 14, 1998 | |
| | C91 | GenBank Accession No. CAA92588 for Similarity to Drosophila potassium channel protein (PIR Acc. No. JH0697); cDNA EST EMBL: D27674 comes from this gene; cDNA EST EMBL: D27673 comes from this gene; cDNA EST EMBL: D32607 comes from this gene; cDNA EST EMBL: D35287 comes from this gene; cDNA EST EMBL: D64971 comes from this gene; cDNA EST EMBL: D68246 comes from this gene; November 23, 1998 | |

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| First Named Inventor | Rory A.J. Curtis |
| Art Unit | 1647 |
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|--------------------|-----------------------|---|----------------|
| /MA/ ↓ | C92 | GenBank Accession No. CAA93875 for Similarity to yeast potassium channel protein; November 23, 1998 | |
| | C93 | GenBank Accession No. CAA93881 for Similarity to potassium channel proteins; November 23, 1998 | |
| | C94 | GenBank Accession No. CAA94204 for Predicted using Genefinder; Similarity to yeast potassium channel protein TOK1; November 23, 1998 | |
| | C95 | GenBank Accession No. CAA95801 for Predicted using Genefinder; weak similarity to yeast potassium channel protein TOK1; November 23, 1998 | |
| | C96 | GenBank Accession No. CAA98271 for Similar to potassium channel protein like; December 18, 1998 | |
| | C97 | GenBank Accession No. CAA98557 for Predicted using Genefinder; Similarity to potassium channel protein; November 23, 1998 | |
| | C98 | GenBank Accession No. CAA98871 for Similar to potassium channel protein; November 23, 1998 | |
| | C99 | GenBank Accession No. CAB01238 Predicted using Genefinder; similarity to worm potassium channel proteins; November 23, 1998 | |
| | C100 | GenBank Accession No. CAB01740 for Similar to potassium channel protein; November 23, 1998 | |
| | | | |

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Sheet 12 of 15

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|------------------------|-------------------|
| Application Number | 10/701,013 |
| Filing Date | 11/04/2003 |
| First Named Inventor | Rory A.J. Curtis |
| Art Unit | 1647 |
| Examiner Name | Allen, Marianne P |
| Attorney Docket Number | 004974.01183 |

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|--------------------|------------------------|---|----------------|
| /MA/ | C101 | GenBank Accession No. CAB02119 for Similar to potassium channel protein like, November 23, 1998 | |
| | C102 | GenBank Accession No. CAB03071 Predicted using Genefinder, similarity to potassium channel proteins; November 23, 1998 | |
| | C103 | GenBank Accession No. CAB03109 Similar to potassium channel protein; November 23, 1998 | |
| | C104 | GenBank Accession No. CAB03914 for C24H11.6, November 23, 1998 | |
| | C105 | GenBank Accession No. CAB04251 for cDNA EST yk240f2.5 comes from this gene; cDNA EST yk354h3.5 comes from this gene; cDNA EST yk354h3.3 comes from this gene; cDNA EST yk240f2.3 comes from this gene, November 23, 1998 | |
| | C106 | GenBank Accession No. CAB04923 for cDNA EST EMBL: D68510 comes from this gene; cDNA EST EMBL: D70321 comes from this gene, November 23, 1998 | |
| | C107 | GenBank Accession No. CAB04928 for cDNA EST yk224a2.3 comes from this gene; cDNA EST yk224a2.5 comes from this gene, November 23, 1998 | |
| | C108 | GenBank Accession No. CAB05769 Similar to potassium channel protein; November 23, 1998 | |
| | C109 | GenBank Accession No. CAB07286 for T28A8.1, November 23, 1998 | |

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/Marianne Allen/

Date
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| First Named Inventor | Rory A.J. Curtis |
| Art Unit | 1647 |
| Examiner Name | Allen, Marianne P |
| Attorney Docket Number | 004974.01183 |

Sheet 13 of 15

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|-----------------------|--------------------------|---|----------------|
| /MA/ | C110 | GenBank Accession No. CAB07375 for F31D4.7, November 23, 1998 | |
| | C111 | GenBank Accession No. CAB07854 for R12G8.2, November 23, 1998 | |
| | C112 | GenBank Accession No. J05677 for RAT guanylyl cyclase A/atrial natriuretic peptide receptor (GC-A) gene, January 15, 1991 | |
| | C113 | GenBank Accession No. L17688 for Human STS UT508B, May 28, 1993 | |
| | C114 | GenBank Accession No. M83796 for Human DNA from cosmid MMDA from Chromosome 19q13.3, April 9, 1992 | |
| | C115 | GenBank Accession No. P34410 for TWK-8 protein, July 15, 1998 | |
| | C116 | GenBank Accession No. P40310 for Outward-rectifier potassium channel TOK1 (Two-domain outward rectifier K ⁺ channel York), November 1, 1997 | |
| | C117 | GenBank Accession No. U31663 for Mus Musculus Chromosome 2 marker um-m12 TC dinucleotide 2 DNA sequence, August 30, 1995 | |
| ↓ | C118 | GenBank Accession No. U31781 for Mus Musculus cGMP-phosphodiesterase beta-subunit gene, May 21, 1996 | |

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Sheet 14 of 15

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| Application Number | 10/701,013 |
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| First Named Inventor | Rory A.J. Curtis |
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|--------------------|-----------------------|---|----------------|
| /MA/ | C119 | GenBank Accession No. U33632 for Human two P-domain K ⁺ channel TWIK-1 mRNA, June 5, 1996 | |
| | C120 | GenBank Accession No. U55321 for Drosophila Melanogaster two P domain potassium channel ORK1 (ORK1) mRNA, October 29, 1998 | |
| | C121 | GenBank Accession No. U73488 for Mus Musculus TREK-1 K ⁺ channel subunit mRNA, January 25, 1997 | |
| | C122 | GenBank Accession No. U76986 for Homo Sapiens two P domain channel subunit (HOKO1) mRNA, January 29, 1998 | |
| | C123 | GenBank Accession No. U90065 for Human potassium channel KCNO1 mRNA, April 1, 1997 | |
| | C124 | GenBank Accession No. Y14406 for Homo Sapiens DNA for 3' IgH locus control region, HS3 enhancer, February 12, 1996 | |
| | C125 | GenBank Accession No. Z55211 for Homo Sapiens CpG island DNA genomic MseI fragment, clone 26g4, FORWARD READ CPG26G4.FT1B, October 17, 1995 | |
| | C126 | GenBank Accession No. Z75539 for Caenorhabditis Elegans cosmid F28C1, December 14, 1998 | |
| | C127 | GenBank Accession No. AC005661 for citb_54_o_2, DNA, November 30, 1999 | |

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| /MA/ | C128 | GenBank Accession No. AC005680 for c1b_43_a_11, DNA; November 5, 1999 | |
| | C129 | GenBank Accession No. Z78198 for Caenorhabditis elegans cosmid F55C5, DNA; January 26, 2000 | |
| | C130 | GenBank Accession No. Z81475 for Caenorhabditis elegans cosmid C24H11, DNA; December 14, 1999 | |
| /MA/ | C131 | GenBank Accession No. Z92813 for Caenorhabditis elegans cosmid T28A8, DNA; December 14, 1999 | |

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